Figure 1A

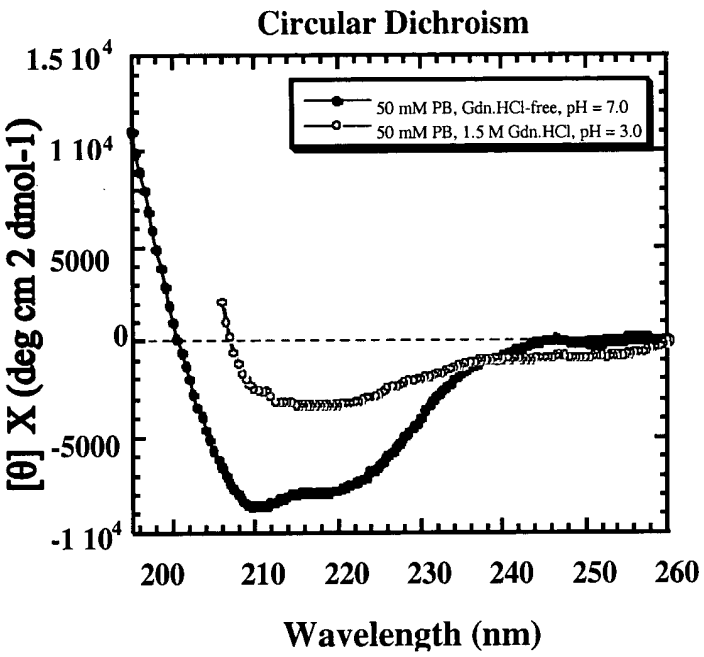


Figure 1B

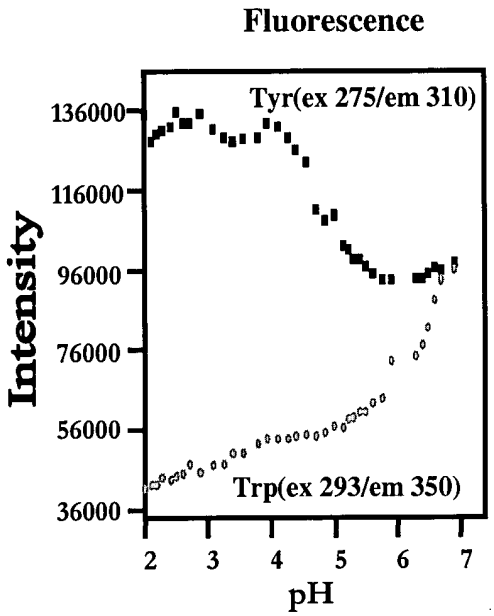
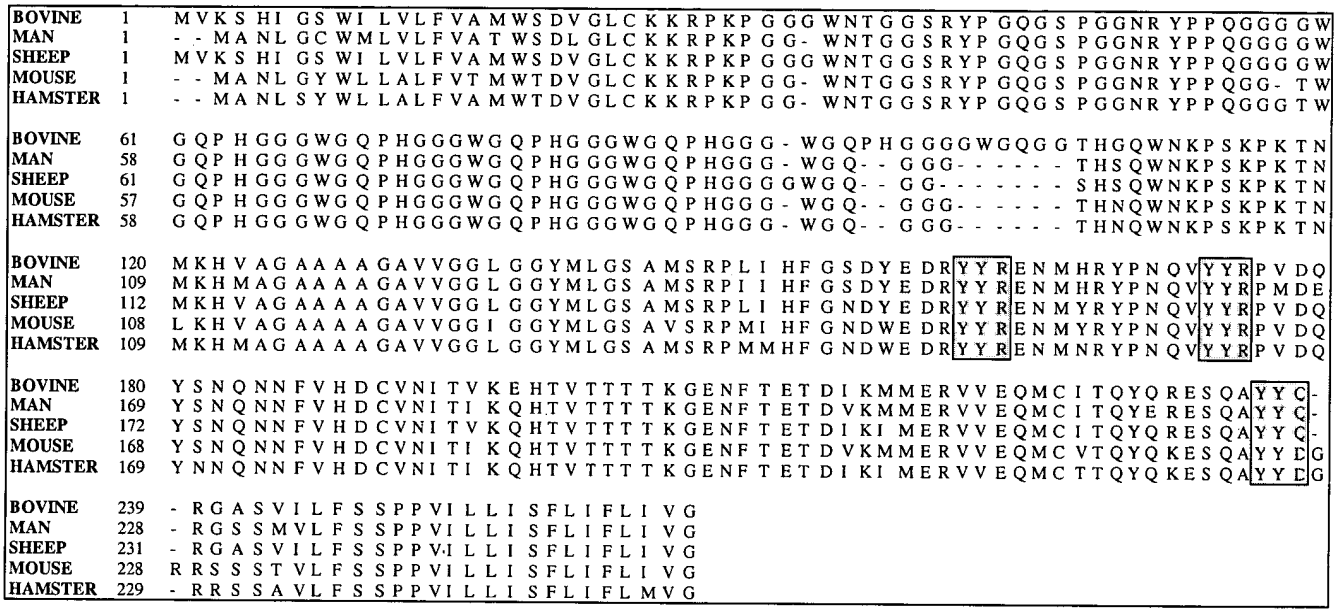


Figure 2



```
BOVINE     1   MVKSHIGSWILVLFVAMWSDVGLCKKRPKPGGGWNTGGSRYPGQGSPGGNRYPPQGGGGW
MAN        1   --MANLGCWMLVLFVATWSDLGLCKKRPKPGG-WNTGGSRYPGQGSPGGNRYPPQGGGGW
SHEEP      1   MVKSHIGSWILVLFVAMWSDVGLCKKRPKPGGGWNTGGSRYPGQGSPGGNRYPPQGGGGW
MOUSE      1   --MANLGYWLLALFVTMWTDVGLCKKRPKPGG-WNTGGSRYPGQGSPGGNRYPPQGG-TW
HAMSTER    1   --MANLSYWLLALFVAMWTDVGLCKKRPKPGG-WNTGGSRYPGQGSPGGNRYPPQGGGTW

BOVINE    61   GQPHGGGWGQPHGGGWGQPHGGGWGQPHGGG-WGQPHGGGGWGQGGTHGQWNKPSKPKTN
MAN       58   GQPHGGGWGQPHGGGWGQPHGGGWGQPHGGG-WGQ--GGG------THSQWNKPSKPKTN
SHEEP     61   GQPHGGGWGQPHGGGWGQPHGGGWGQPHGGGWGQPHGGGGWGQ--GG-------SHSQWNKPSKPKTN
MOUSE     57   GQPHGGGWGQPHGGGWGQPHGGGWGQPHGGGWGQPHGGG-WGQ--GGG------THNQWNKPSKPKTN
HAMSTER   58   GQPHGGGWGQPHGGGWGQPHGGGWGQPHGGG-WGQ--GGG------THNQWNKPSKPKTN

BOVINE   120   MKHVAGAAAAGAVVGGLGGYMLGSAMSRPLIHFGSDYEDRYYRENMHRYPNQVYYRPVDQ
MAN      109   MKHMAGAAAAGAVVGGLGGYMLGSAMSRPIIHFGSDYEDRYYRENMHRYPNQVYYRPMDE
SHEEP    112   MKHVAGAAAAGAVVGGLGGYMLGSAMSRPLIHFGNDYEDRYYRENMYRYPNQVYYRPVDQ
MOUSE    108   LKHVAGAAAAGAVVGGIGGYMLGSAVSRPMIHFGNDWEDRYYRENMYRYPNQVYYRPVDQ
HAMSTER  109   MKHMAGAAAAGAVVGGLGGYMLGSAMSRPMMHFGNDWEDRYYRENMNRYPNQVYYRPVDQ

BOVINE   180   YSNQNNFVHDCVNITVKEHTVTTTTKGENFTETDIKMMERVVEQMCITQYQRESQAYYQ-
MAN      169   YSNQNNFVHDCVNITIKQHTVTTTTKGENFTETDVKMMERVVEQMCITQYERESQAYYQ-
SHEEP    172   YSNQNNFVHDCVNITVKQHTVTTTTKGENFTETDIKIMERVVEQMCITQYQRESQAYYQ-
MOUSE    168   YSNQNNFVHDCVNITIKQHTVTTTTKGENFTETDVKMMERVVEQMCVTQYQKESQAYYDG
HAMSTER  169   YNNQNNFVHDCVNITIKQHTVTTTTKGENFTETDIKIMERVVEQMCTTQYQKESQAYYDG

BOVINE   239   -RGASVILFSSPPVILLISFLIFLIVG
MAN      228   -RGSSMVLFSSPPVILLISFLIFLIVG
SHEEP    231   -RGASVILFSSPPVILLISFLIFLIVG
MOUSE    228   RRSSSTVLFSSPPVILLISFLIFLIVG
HAMSTER  229   -RRSSAVLFSSPPVILLISFLIFLMVG
```

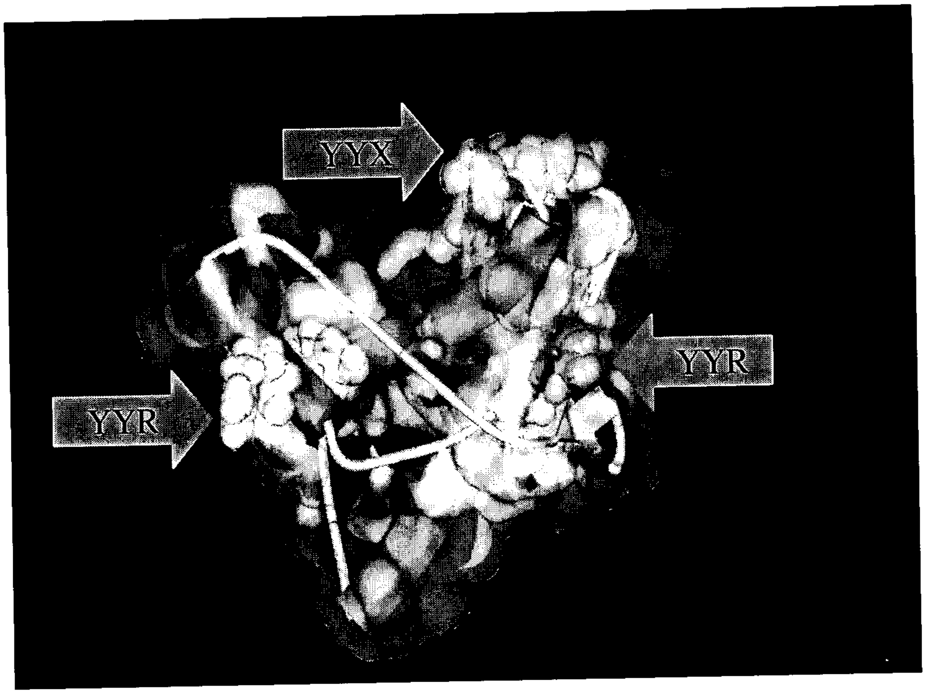


Figure 3

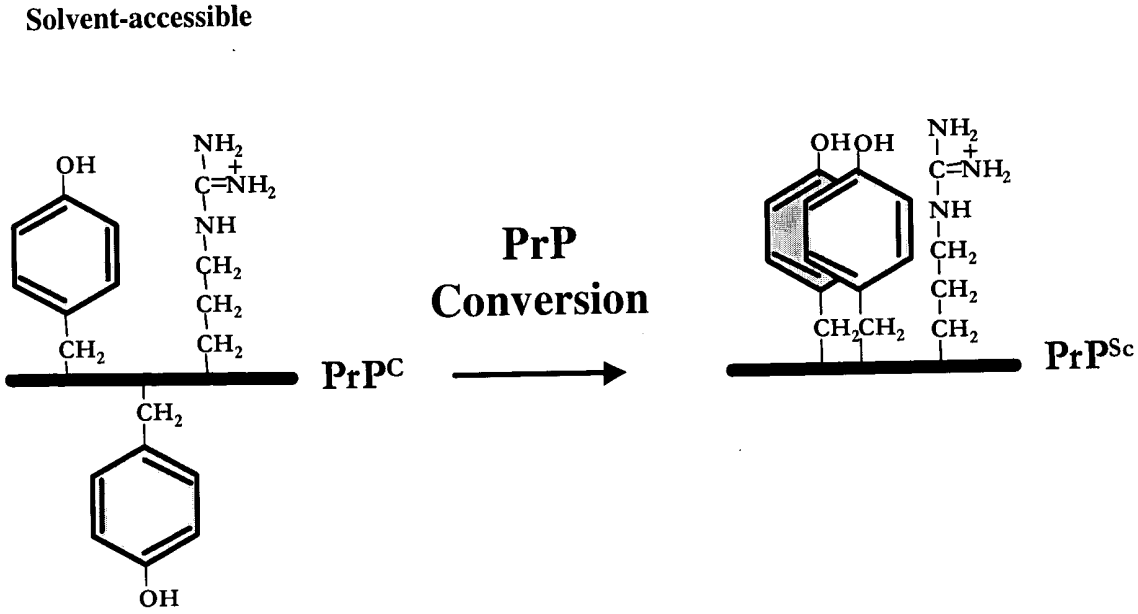

Solvent-accessible
PrP
Conversion
PrP^C
PrP^Sc
Solvent-inaccessible

Figure 4

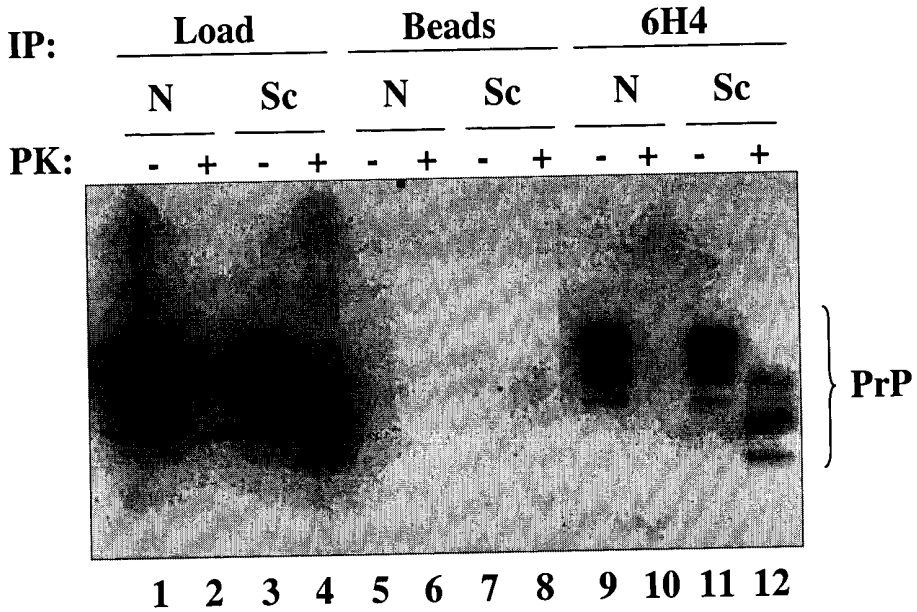


Figure 5

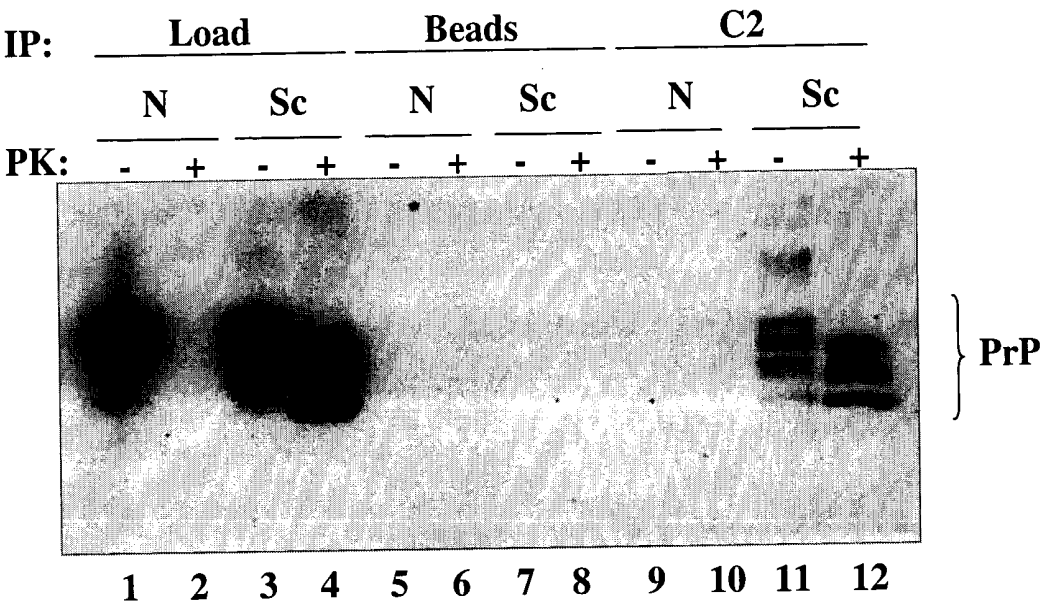


Figure 6

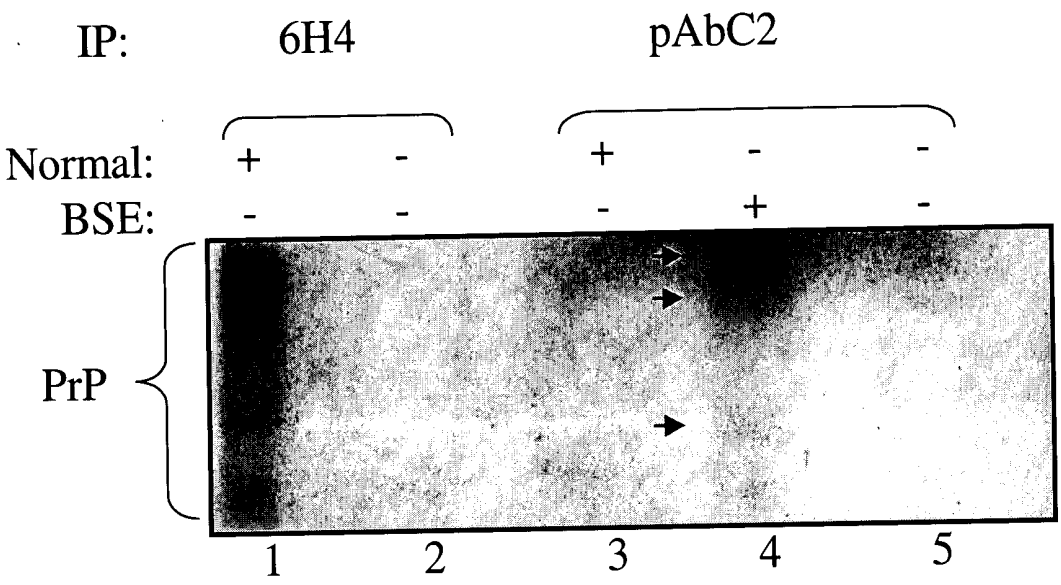


Figure 7

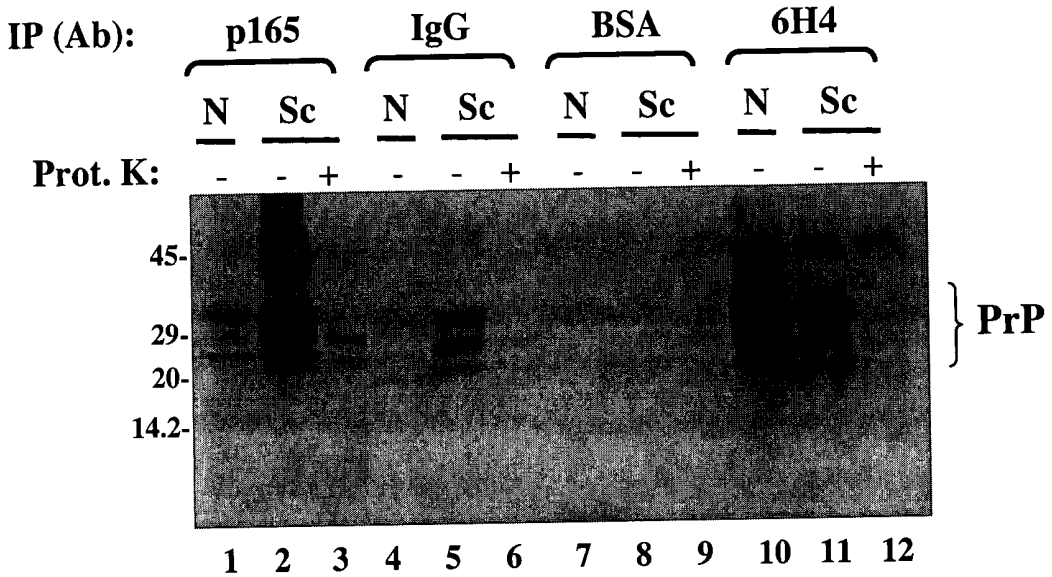

IP (Ab):
p165
IgG
BSA
6H4
N
Sc
N
Sc
N
Sc
N
Sc
Prot. K:
- - + - - + - - + - - +
45-
29-
20-
14.2-
PrP
1 2 3 4 5 6 7 8 9 10 11 12

Figure 8

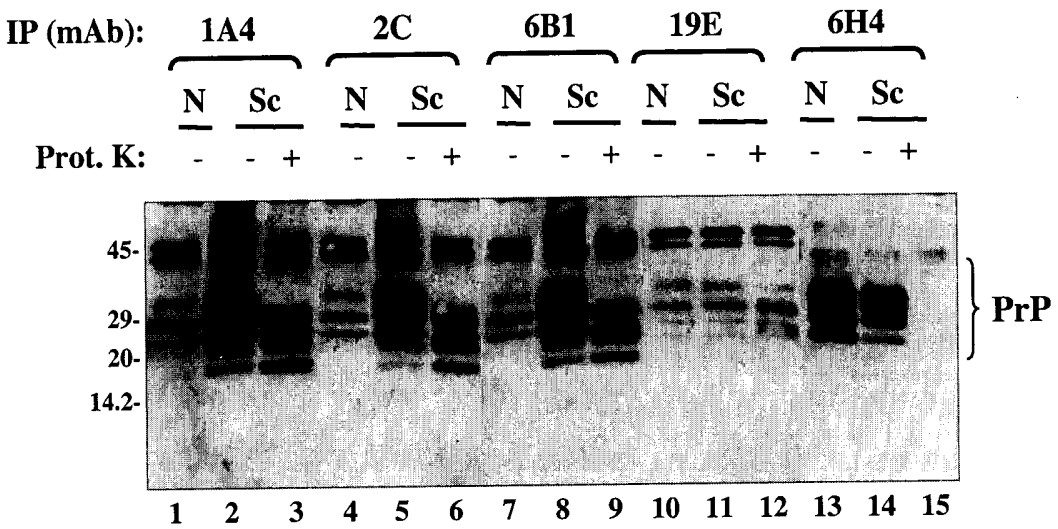

IP (mAb):
1A4
2C
6B1
19E
6H4
N
Sc
Prot. K:
- + - + - + - + - + - + - + - + - +
45-
29-
20-
14.2-
PrP
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15

Figure 9

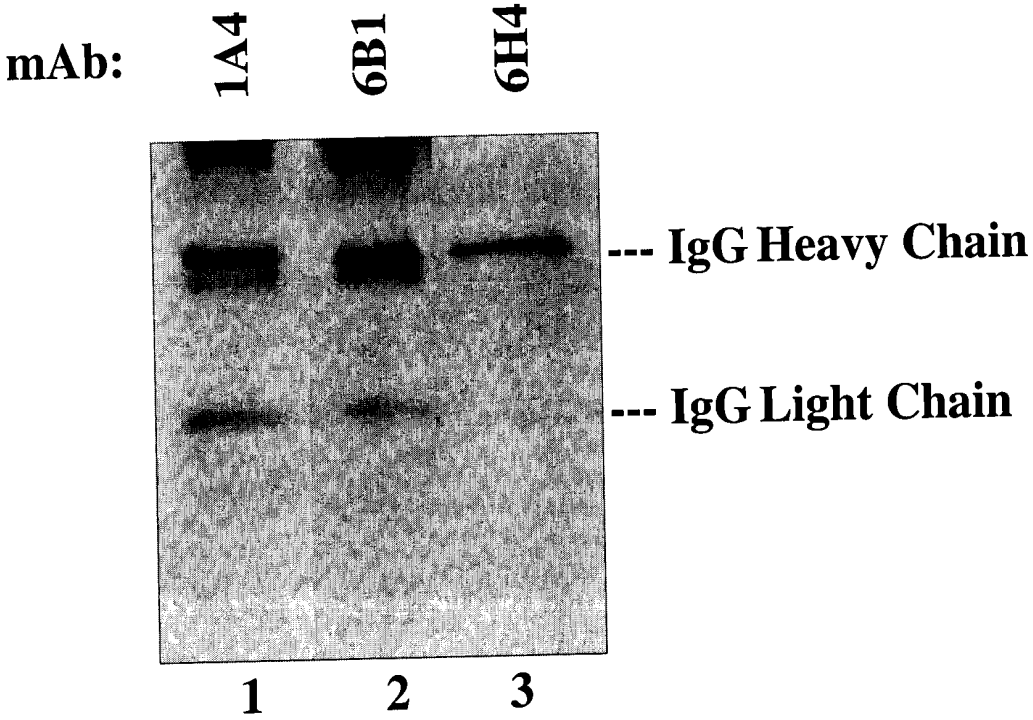


Figure 10

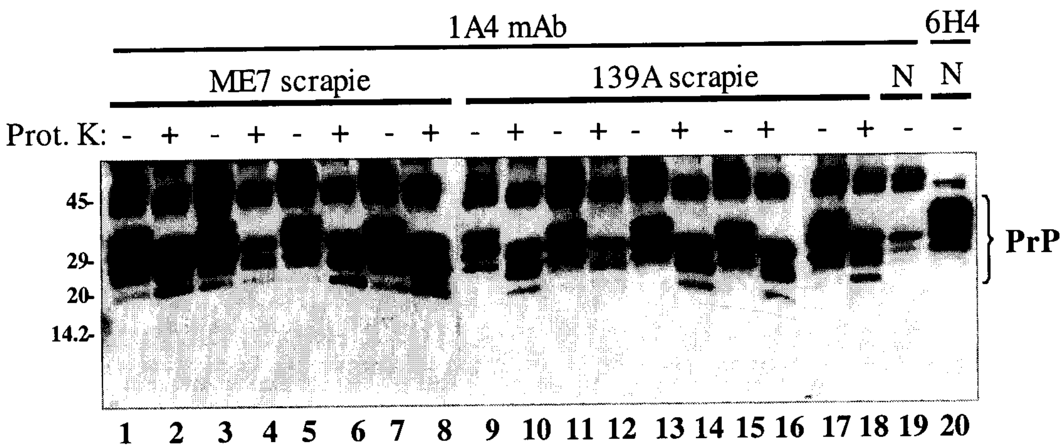

1A4 mAb
6H4
ME7 scrapie
139A scrapie
N
N
Prot. K: - + - + - + - + - + - + - + - + - + - -
45-
29-
20-
14.2-
PrP
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20

Figure 11

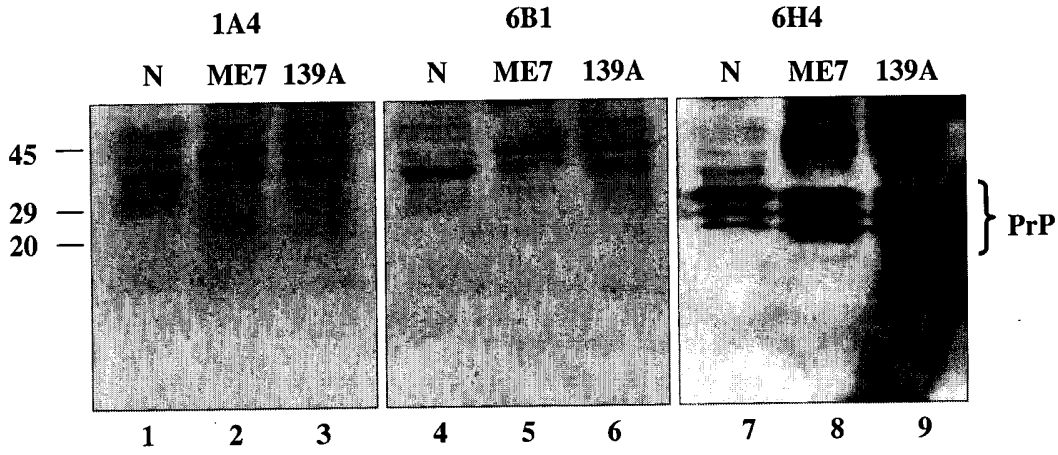
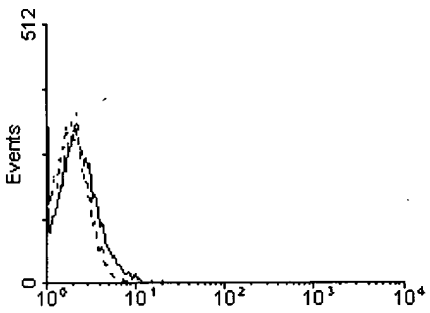
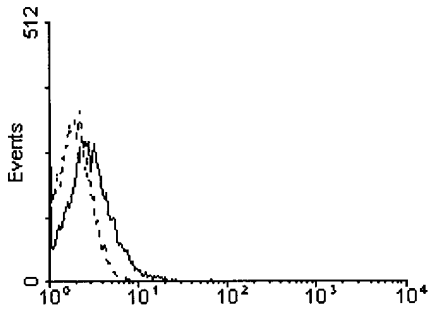

1A4
6B1
6H4
N
ME7
139A
N
ME7
139A
N
ME7
139A
45
29
20
PrP
1
2
3
4
5
6
7
8
9

Figure 12

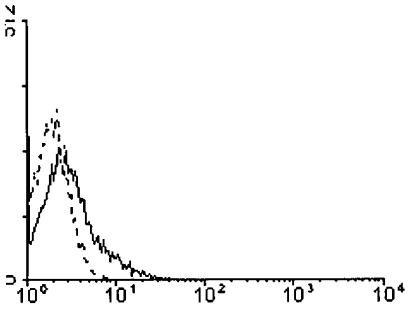
Figure 13



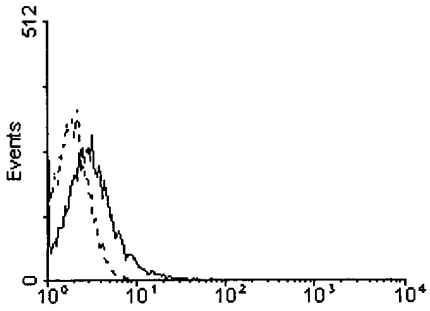
1A4-FITC



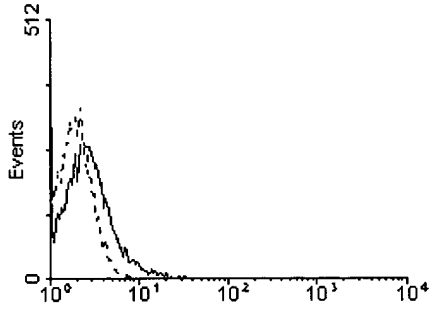
2B5-FITC



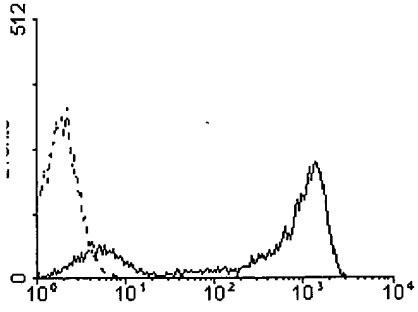
6B1-FITC



17B-FITC



18B-FITC



GAMIg-FITC

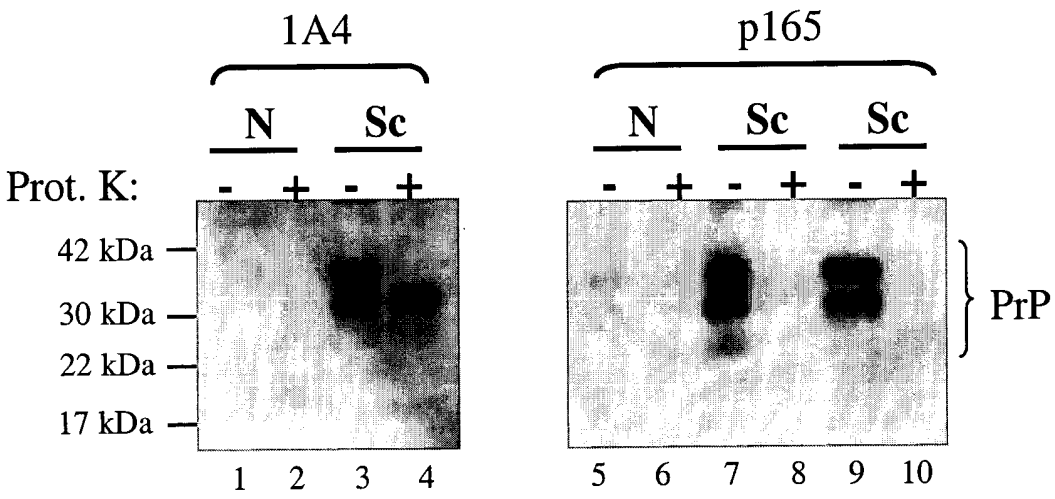

1A4
p165
N
Sc
N
Sc
Sc
Prot. K:
- + - + - + - +
42 kDa
30 kDa
22 kDa
17 kDa
PrP
1 2 3 4 5 6 7 8 9 10

Figure 14

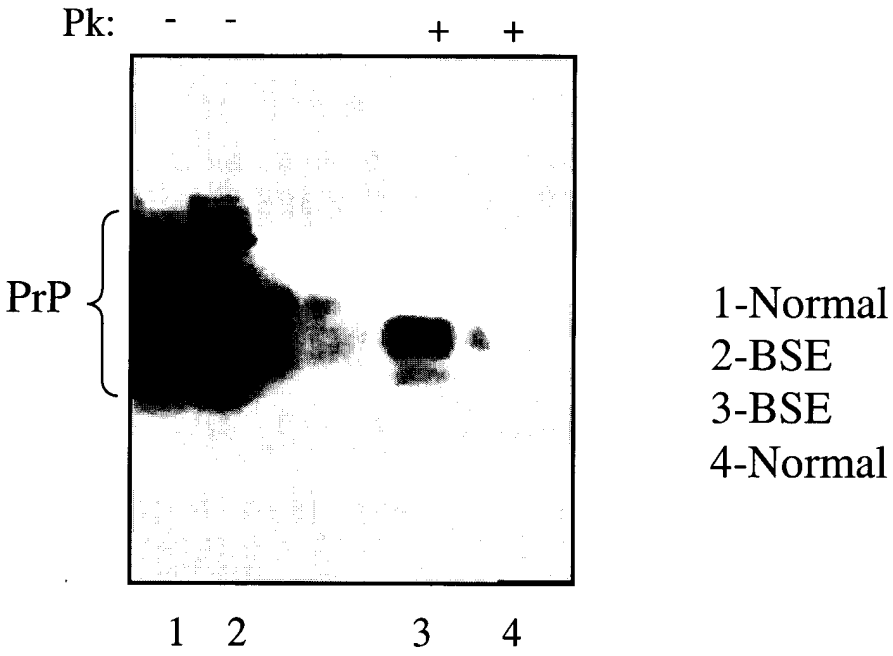
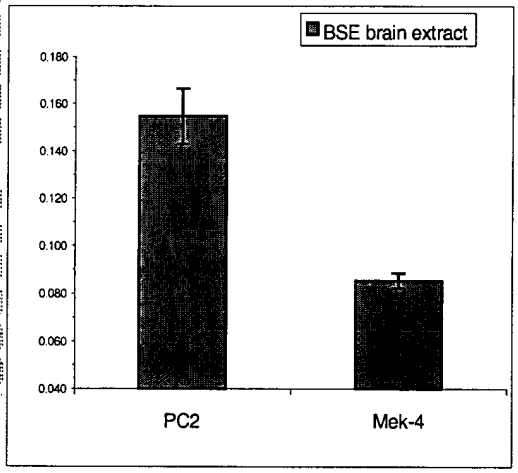


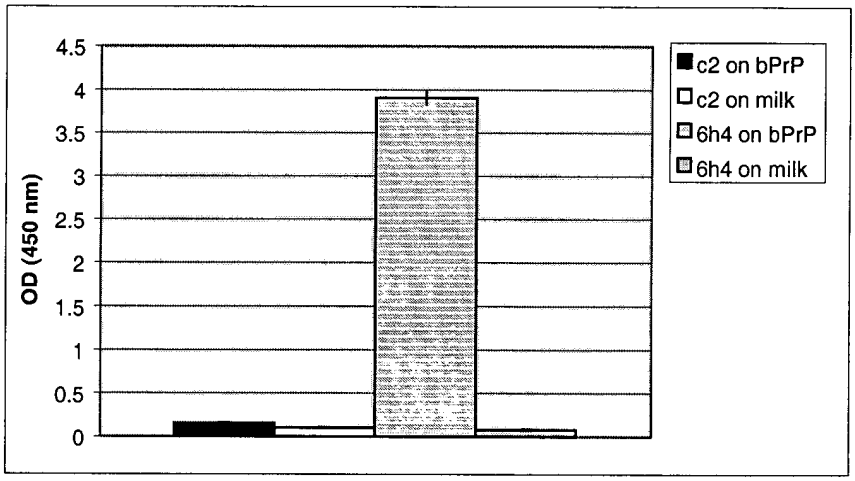
Figure 15

Figure 16

A


BSE brain extract
0.180
0.160
0.140
0.120
0.100
0.080
0.060
0.040
PC2
Mek-4

B


OD (450 nm)
4.5
4
3.5
3
2.5
2
1.5
1
0.5
0
c2 on bPrP
c2 on milk
6h4 on bPrP
6h4 on milk